

SEQUENCE LISTING

<110> Blenis, John
Murphy, Leon O.

<120> Inhibitors of the Map Kinase Pathway

<130> 00742/074002

<140> US 10/563,345

<141> 2006-01-03

<150> PCT/US04/21514

<151> 2004-07-02

<150> US 60/484,761

<151> 2003-07-03

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Cys Thr Pro Ser Cys Thr Thr Tyr Thr Ser Ser Phe Val Phe Thr Tyr

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 Gly Gly Phe Tyr Gly Glu Glu Pro Leu His Thr Pro Ile Val Val Thr
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 Ser Thr Pro Ala Ile Thr Pro Gly Thr Ser Asn Leu Val Phe Thr Tyr
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 Pro Ser Val Leu Glu Gln
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 Leu Leu Ala Ser Pro Asp Leu Gly Leu Leu Lys Leu Ala Ser Pro Glu
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 Thr Ser Thr Gln Phe Leu Tyr Pro Lys Val
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 Gly Gln Gly Ser Asp Thr Gly Ala Ser Leu Lys Leu Ala Ser Ser Glu
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Pro Thr Pro Pro Gly Gln Tyr Phe Tyr Pro Arg Gly
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Ala His Pro Ala Ala Glu Cys Val Asp Pro Ala Val Val Phe Pro Phe
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Pro Val Asn Lys
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Pro Met Ile Pro Asp Tyr Leu Phe Pro Gln Gln
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Pro Arg Gly Gly Pro Gln Pro Leu Pro Pro Ala Pro Thr Ser Val Pro
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Lys Pro Ser Thr Phe Ala Tyr Pro Ala Pro Leu Glu Val Pro Lys
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Ser Lys Ile Ser Trp Glu Phe Pro Glu Ser Ser Ser Ser Glu Glu
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Pro	Leu	Leu	Thr	Tyr	Arg	Phe	Pro	Pro	Lys	Phe	Thr	Leu	Lys	Ala
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Ser	Val	Pro	Asp	Phe	Arg	Phe	Pro	Met	Ala	Asp	Gly	His	Thr	Asp
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<400> 147

Val	Asn	Arg	Arg	Phe	His	Phe	Pro	Ile	Tyr	Tyr	Leu	Met	Ala	Asn
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<400> 148

Lys	Asn	Arg	Lys	Phe	His	Phe	Pro	Phe	Tyr	Tyr	Leu	Leu	Ala	Asn
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<400> 149

Ser	Ile	His	Ser	Phe	Gly	Phe	Pro	Asn	Phe	Val	Lys	Lys	Ile	Asp
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<400> 151

Arg	Ala	Ser	Leu	Tyr	Ser	Phe	Pro	Glu	Pro	Glu	Ala	Ala	Gln	Pro
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<400> 152

Gly	Asn	Trp	Gln	Tyr	Phe	Phe	Pro	Val	Ile	Phe	Ser	Lys	Ala	Ser
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Pro Leu Asn Tyr Tyr Asn Phe Pro Lys Ser Cys Cys Thr Ser Val
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Lys Pro Cys Cys Tyr Ile Phe Pro Gly Gly Arg Gly Asp Ser Ala
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Lys Asn Thr Ser Phe Ala Tyr Pro Ala Ile Arg Tyr Leu Leu Tyr
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Tyr Met Ser Ile Tyr Ser Phe Pro Met Asp Glu Pro Ile Ser Ser

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 Thr Ala Ala Ala Phe Asp Phe Pro Thr Ser Phe Gln Leu Thr Glu
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 Thr Lys His Pro Phe Ser Phe Pro Leu Glu Asn Gln Ala Arg Glu
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Tyr	Val	Gly	Arg	Phe	Lys	Phe	Pro	Glu	Glu	Arg	Ala	Ala	Glu	Val
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Gly	Tyr	Val	Asn	Phe	Arg	Phe	Pro	Ala	Asp	Ala	Glu	Trp	Ala	Leu
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Ser	Glu	Leu	Glu	Phe	Gln	Phe	Pro	Asp	Asp	Lys	Asp	Pro	Val	Trp
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Asn	Leu	Ile	Gly	Phe	Gly	Tyr	Pro	Ala	Tyr	Ile	Ser	Ile	Lys	Ala
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<400> 205

Lys	Gly	Ser	Ala	Phe	Thr	Phe	Pro	Ser	Gln	Gln	Ser	Pro	Arg	Asn
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<213> Homo sapiens

<400> 206

Lys	Lys	Gly	Glu	Tyr	Val	Phe	Pro	Ala	Lys	Lys	Lys	Leu	Gln	Glu
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<213> Homo sapiens

<400> 207

Lys	Asn	Val	Cys	Phe	Ile	Phe	Pro	Gln	Phe	Leu	Tyr	Gln	Phe	Phe
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<400> 208

Val	Tyr	Asn	Ser	Phe	Gly	Phe	Pro	Gly	Gln	Ala	Ala	Arg	Tyr	Pro
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<213> Homo sapiens

<400> 209

Val	Phe	Glu	Thr	Phe	Met	Phe	Pro	Cys	Gln	His	Cys	Glu	Arg	Lys
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Val Phe Val Gly Phe Leu Phe Pro Trp Ala Leu Met Leu Leu Ser
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Phe Leu Val Gly Phe Leu Phe Pro Ile Cys Leu Leu Leu Ala Ser
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Ala Phe Ile Thr Phe Met Phe Pro Glu His Ala Val Lys Ala Tyr
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Gln Glu Ala Arg Phe Asp Phe Pro Gly Ser Ala Gly Val Asp Arg
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Phe Asn Asp Tyr Phe Glu Phe Pro Arg Glu Leu Asp Met Glu Pro

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Thr Ser Thr Asp Phe Ser Phe Pro Asp Val Asn Glu Lys Asp Ala
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His Tyr Ser Thr Phe Ala Tyr Pro Pro Thr Glu Val Thr Ser His
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Leu Tyr Gly Gly Phe Pro Phe Pro Leu Glu Met Glu Asn Lys Arg
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Asn Asn Gln Leu Phe Arg Phe Pro Ala Thr Ser Pro Leu Lys Thr
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Ile Ile Pro Gly Phe Pro Tyr Pro Thr Ala Ala Thr Thr Ala Ala
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Val	Ser	Pro	Arg	Phe	Ser	Phe	Pro	Gly	Thr	Thr	Gly	Ser	Leu	Ala
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Glu	Pro	Glu	Asp	Phe	Ala	Phe	Pro	Ser	Thr	Ala	Pro	Ser	Pro	Gln
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Ala	Pro	Gln	Pro	Tyr	Thr	Phe	Pro	Ala	Ser	Leu	Ser	Thr	Ile	Asn
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Thr	Ala	Leu	Gln	Phe	Thr	Tyr	Pro	Leu	Phe	Thr	Thr	Asn	Ala	Cys
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<400> 282

Pro	Pro	Glu	Asn	Tyr	Asp	Phe	Pro	Val	Val	Ile	Val	Lys	Gln	Glu
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<400> 283

Gly	Gly	Ala	Ala	Phe	Ile	Phe	Pro	Asn	Thr	Ser	Val	Tyr	Pro	Glu
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Gly	Ser	His	Leu	Phe	Gly	Phe	Pro	Pro	Thr	Pro	Pro	Lys	Glu	Val
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His	Ala	Gln	Pro	Phe	Asp	Phe	Pro	Asp	Asp	Asn	Gln	Asn	Ser	Lys
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Asn	Val	Pro	Asp	Phe	Asp	Phe	Pro	Pro	Glu	Phe	Tyr	Glu	His	Ala
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Phe	Pro	Val	Arg	Phe	Pro	Tyr	Pro	Cys	Thr	Gln	Thr	Glu	Leu	Ala
1				5					10					15

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Gly	Leu	Glu	Thr	Tyr	Gly	Phe	Pro	Ser	Val	Thr	Leu	Ile	Ser	Cys
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<400> 318

Ser	Thr	Asn	Pro	Phe	Asp	Phe	Pro	Phe	Val	Ser	Gln	Gly	Glu	Val
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<213> Homo sapiens

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Phe	Pro	Gln	Gly	Tyr	Gln	Phe	Pro	Ser	Met	Glu	Gln	Leu	Ala	Ala
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Pro	Met	Asp	Arg	Phe	Ser	Phe	Pro	Asp	Gly	Lys	Glu	Ser	Asp	Thr
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Asp	Val	Ala	Asp	Tyr	Val	Phe	Pro	Ala	Gln	Gly	Asp	Asn	Ser	Phe
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Asp	Thr	Ala	Asp	Tyr	Thr	Phe	Pro	Leu	Gln	Gly	Asn	Ser	Phe	Phe
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<400> 323

Val	Phe	Val	Gly	Phe	Leu	Phe	Pro	Trp	Ala	Leu	Met	Leu	Leu	Ser
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Asn Val Ala Ser Phe Leu Tyr Pro Asn Leu Gly Gly Ser Trp Arg
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Ala Pro Tyr His Phe Ser Phe Pro Leu Asp Phe Leu Val Lys Ser
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Lys Pro Asp Leu Phe Thr Tyr Pro Cys Glu Met Ser Leu Ile Ser
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 Pro Tyr Gly Val Phe Ile Phe Pro Asn Lys Thr Arg Pro Leu Ser
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 Asp Gly Leu Lys Tyr Leu Phe Pro Gly Glu Cys Gln Tyr Val Leu
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Phe	Pro	Val	Arg	Phe	Pro	Tyr	Pro	Cys	Thr	Gln	Thr	Glu	Leu	Ala
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Gln	Phe	His	Ser	Phe	Ile	Tyr	Pro	Tyr	Met	Ala	Asn	Gly	Ser	Leu
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Met	Glu	Glu	Glu	Phe	Asn	Tyr	Pro	Leu	Asp	Asn	Val	His	Leu	Leu
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Gln	Cys	Gly	Leu	Phe	Ser	Phe	Pro	Cys	Lys	Asn	Gly	Arg	Cys	Val
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Val	Pro	Cys	Ser	Phe	Ser	Tyr	Pro	Trp	Arg	Ser	Trp	Tyr	Ser	Ser
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Gly	Gly	Gly	Ala	Phe	Met	Phe	Pro	Tyr	Phe	Ile	Met	Leu	Ile	Phe
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Ile	Gly	Pro	Leu	Phe	Phe	Phe	Pro	Leu	Leu	Tyr	Met	Ile	Phe	Gln
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Pro	Arg	Ser	Ser	Phe	Ala	Phe	Pro	Pro	Ser	Leu	Ala	Lys	Ala	Gly
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Glu	Tyr	Gly	Ser	Tyr	Arg	Phe	Pro	Pro	Trp	Ala	Glu	Leu	Leu	Gly
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Ser	Leu	Leu	Met	Phe	Gly	Phe	Pro	Gln	Ser	Leu	Pro	Pro	His	Ser
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Phe	Asp	Glu	Leu	Phe	Gln	Phe	Pro	Val	Val	Tyr	Asp	Gln	Leu	Ser
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Phe	Ser	Val	Asp	Tyr	Asn	Phe	Pro	Lys	Lys	Ala	Leu	Val	Val	Thr
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Gly	Asp	Glu	Glu	Phe	Leu	Phe	Pro	Asp	Lys	Lys	Asp	Arg	Gln	Asn
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Ala	Trp	Lys	Tyr	Tyr	Lys	Phe	Pro	Lys	Lys	Lys	Asp	Val	Glu	Phe
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<400> 493

Lys	Thr	Cys	Pro	Phe	Phe	Phe	Pro	Phe	Asp	Thr	Arg	Gln	Met	Leu
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<400> 494

Tyr	Thr	Val	Arg	Phe	Thr	Phe	Pro	Asp	Pro	Pro	Pro	Leu	Ser	Pro
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Gly Phe Gly Ser Phe Arg Phe Pro Ser Gly Asn Gln Gly Gly Ala
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<400> 496
Phe Thr Met Asp Phe Cys Phe Pro Phe Ser Pro Leu Gln Ala Phe
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Tyr Ile Leu Asp Phe Gln Tyr Pro Phe Ser Ala Val Gln Ala Phe
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His Ser Lys Thr Tyr His Phe Pro Lys Thr Val Thr Gln Ser Leu
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Glu Thr Cys Ser Phe Cys Phe Pro Glu Cys Arg Ala Pro Thr Gln
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Tyr Arg Ile Arg Phe Tyr Phe Pro Arg Trp Tyr Cys Ser Gly Ser

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 Cys Trp Ala Asp Phe Gly Phe Pro Trp Gly Pro Arg Ala Tyr Leu
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 Pro Tyr Gly Val Phe Ile Phe Pro Asn Lys Thr Arg Pro Leu Ser
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 Lys Ser Leu Cys Phe Gln Tyr Pro Pro Val Tyr Val Gly Lys Ile
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 Cys Gly Lys Gly Phe Asp Phe Pro Gly Ser Ala Arg Ile His Glu
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 Asp Gly Gly Arg Phe Lys Phe Pro Asp Gly Val Gln Asn Trp Pro
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 Cys Ala His Trp Phe Gln Tyr Pro Phe Asp Pro Ser Phe Asp Asp
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Glu Leu Gln Gln Phe Gly Phe Pro Ala Pro Gln Thr Gly Asp Pro
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Ser Tyr Ala Lys Phe Leu Tyr Pro Thr Asn Ala Leu Val Thr His
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Gly Ala Lys Ala Phe Tyr Tyr Pro Lys Glu Ala Gly Val Pro Phe
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<400> 546

Arg	Leu	Ser	Ala	Phe	Val	Phe	Pro	Gly	Glu	Leu	Leu	Leu	Arg	Leu
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Thr	Thr	Trp	Val	Phe	Ser	Phe	Pro	Phe	Cys	Gly	Pro	Asn	Glu	Ile
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Thr	Thr	Trp	Leu	Phe	Ser	Phe	Pro	Phe	Cys	Gly	Thr	Asn	Lys	Val
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<400> 549

Ser	Tyr	Ile	Tyr	Phe	Leu	Phe	Pro	Pro	Leu	Met	Asn	Pro	Val	Ile
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<212> PRT

<213> Homo sapiens

<400> 550

Ser	Tyr	Val	His	Phe	Leu	Phe	Pro	Pro	Phe	Val	Asn	Pro	Ile	Ile
1				5					10					15

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<212> PRT

<213> Homo sapiens

<400> 551

Phe	Thr	Thr	Leu	Phe	Pro	Phe	Pro	Phe	Val	Val	Lys	Arg	Leu	Pro
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Ala Leu Val Ile Phe Leu Phe Pro Leu Phe Ile Thr Val Leu Ser
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Asn Cys Thr Met Phe Arg Phe Pro Leu Arg Asn Ala Glu Met Ala
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 Ser Gly Lys Val Phe Val Phe Pro Arg Glu Ser Val Thr Asp His
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 Thr Met Gly Asn Tyr Val Phe Pro Lys Trp Gly Gln Gly Val Gly
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 Lys Lys Asp Arg Tyr Val Phe Pro Leu Pro Tyr Leu Gly Asp Thr
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 Arg Tyr Pro Asn Tyr Met Phe Pro Ser Gln Gly Ile Thr Pro Gln
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 Phe Ser Pro Arg Phe Pro Phe Pro Thr Val Pro Pro Ala Pro Gly
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Phe Phe Val Ile Phe Ser Phe Pro Ile Ala Ser Lys Asp Cys Ile
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Pro Tyr Pro Ala Phe Pro Phe Pro Tyr Leu Asp Thr Phe Met Thr
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Ser Asp Ile Ser Phe Glu Tyr Pro Glu Asp Ile Gly Asn Leu Ser
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